

=====

Sequence Listing was accepted.

See attached Validation Report.

If you need help call the Patent Electronic Business Center at (866) 217-9197 (toll free).

Reviewer: Anne Corrigan

Timestamp: [year=2008; month=11; day=2; hr=15; min=50; sec=48; ms=205;]

=====

Application No: 10579104 Version No: 2.0

Input Set:

Output Set:

Started: 2008-09-29 16:16:48.125
Finished: 2008-09-29 16:16:51.056
Elapsed: 0 hr(s) 0 min(s) 2 sec(s) 931 ms
Total Warnings: 44
Total Errors: 14
No. of SeqIDs Defined: 44
Actual SeqID Count: 44

Error code	Error Description
W 213	Artificial or Unknown found in <213> in SEQ ID (1)
W 213	Artificial or Unknown found in <213> in SEQ ID (2)
W 213	Artificial or Unknown found in <213> in SEQ ID (3)
W 213	Artificial or Unknown found in <213> in SEQ ID (4)
W 213	Artificial or Unknown found in <213> in SEQ ID (5)
W 213	Artificial or Unknown found in <213> in SEQ ID (6)
W 213	Artificial or Unknown found in <213> in SEQ ID (7)
W 213	Artificial or Unknown found in <213> in SEQ ID (8)
E 257	Invalid sequence data feature in <221> in SEQ ID (8)
W 213	Artificial or Unknown found in <213> in SEQ ID (9)
E 257	Invalid sequence data feature in <221> in SEQ ID (9)
W 213	Artificial or Unknown found in <213> in SEQ ID (10)
E 257	Invalid sequence data feature in <221> in SEQ ID (10)
W 213	Artificial or Unknown found in <213> in SEQ ID (11)
E 257	Invalid sequence data feature in <221> in SEQ ID (11)
W 213	Artificial or Unknown found in <213> in SEQ ID (12)
E 257	Invalid sequence data feature in <221> in SEQ ID (12)
W 213	Artificial or Unknown found in <213> in SEQ ID (13)
E 257	Invalid sequence data feature in <221> in SEQ ID (13)
W 213	Artificial or Unknown found in <213> in SEQ ID (14)

Input Set:

Output Set:

Started: 2008-09-29 16:16:48.125
Finished: 2008-09-29 16:16:51.056
Elapsed: 0 hr(s) 0 min(s) 2 sec(s) 931 ms
Total Warnings: 44
Total Errors: 14
No. of SeqIDs Defined: 44
Actual SeqID Count: 44

Error code	Error Description
E 257	Invalid sequence data feature in <221> in SEQ ID (14)
W 213	Artificial or Unknown found in <213> in SEQ ID (15)
E 257	Invalid sequence data feature in <221> in SEQ ID (15)
W 213	Artificial or Unknown found in <213> in SEQ ID (16)
E 257	Invalid sequence data feature in <221> in SEQ ID (16)
W 213	Artificial or Unknown found in <213> in SEQ ID (17)
E 257	Invalid sequence data feature in <221> in SEQ ID (17)
W 213	Artificial or Unknown found in <213> in SEQ ID (18)
E 257	Invalid sequence data feature in <221> in SEQ ID (18)
W 213	Artificial or Unknown found in <213> in SEQ ID (19)
E 257	Invalid sequence data feature in <221> in SEQ ID (19)
W 213	Artificial or Unknown found in <213> in SEQ ID (20) This error has occurred more than 20 times, will not be displayed
E 257	Invalid sequence data feature in <221> in SEQ ID (20)
E 257	Invalid sequence data feature in <221> in SEQ ID (21)

SEQUENCE LISTING

<110> POLYPHOR LTD.
Universitaet Zuerich
<120> Template fixed beta-hairpin mimetics and their use in phage
display

<130> 753-65 PCT-US

<140> 10579104
<141> 2008-09-29

<150> PCT/EP 03/12783
<151> 2003-11-15

<160> 44

<170> PatentIn version 3.5

<210> 1
<211> 4
<212> PRT
<213> Artificial Sequence

<220>

<223> Key sequence known to occur in Platelet-Derived Growth Factor
(PDGF), see Ross, R.; Raines, E. W.; Bowden-Pope, D.F.; Cell,
1986, 46, 155-159.

<400> 1

Val Arg Lys Lys
1

<210> 2
<211> 4
<212> PRT
<213> Artificial Sequence

<220>

<223> Key sequence known to occur in Vasointestinal Peptide (VIP)
showing neuroprotective properties against beta-amyloid
neurotoxicity, see Proc. Natl. Am. Soc. USA, 1996, 96, 4143-4148.

<400> 2

Lys Lys Tyr Leu
1

<210> 3
<211> 4
<212> PRT
<213> Artificial Sequence

<220>
<223> Key sequence known to occur in integrin alpha.sub4 beta.sub1, see
Europ. J. Biol., 1996, 242, 352-362 and Int. J. Pept. Prot. Res.,
1996, 47, 427-436.

<400> 3

Trp Leu Asp Val

1

<210> 4
<211> 5
<212> PRT
<213> Artificial Sequence

<220>
<223> Key sequence known to occur in Factor Xa inhibitors, see Al
Obeidis, F.; Ostrem, J. A.; Drug Discovery Today, 1998, 3,
223-231.

<400> 4

Tyr Ile Arg Leu Pro

1 5

<210> 5
<211> 5
<212> PRT
<213> Artificial Sequence

<220>
<223> Key sequence known to occur in laminine, see EMBO. J., 1984, 3,
1463.

<400> 5

Tyr Ile Gly Ser Arg

1 5

<210> 6
<211> 5
<212> PRT
<213> Artificial Sequence

<220>
<223> Key sequence known to occur in important physiologically active
peptides, see Cell, 1987, 88, 989.

<400> 6

Ile Lys Val Ala Val

1 5

<210> 7
<211> 6
<212> PRT
<213> Artificial Sequence

<220>
<223> Key sequence known to occur in important physiologically active peptides, see J. Biol. Chem., 1998, 273, 11001-11006 and 11007-11011.

<220>
<221> misc_feature
<222> (4)..(5)
<223> Xaa can be any naturally occurring amino acid

<400> 7

Pro Pro Arg Xaa Xaa Trp
1 5

<210> 8
<211> 10
<212> PRT
<213> Artificial Sequence

<220>
<223> Hairpin mimetic derived from the general formula Cys-Z-Cys wherein the alpha amino group of the first amino acid is acetylated and wherein Z consists of 8 amino acids.

<220>
<221> DISULFID
<222> (1)..(10)

<220>
<221> MOD_RES
<222> (1)..(1)
<223> ACETYLATION

<400> 8

Cys Lys Trp Phe Leu Ala His Tyr Ala Cys
1 5 10

<210> 9
<211> 14
<212> PRT
<213> Artificial Sequence

<220>
<223> Hairpin mimetic derived from the general formula R1-Cys-Z-Cys-R2 wherein the alpha amino group of the first amino acid is

acetylated, wherein Z consists of 8 amino acids, and wherein both R1 and R2 consist of 2 amino acids.

<220>
<221> MOD_RES
<222> (1)..(1)
<223> ACETYLATION

<220>
<221> DISULFID
<222> (3)..(12)

<400> 9

Glu Thr Cys Lys Trp Phe Leu Ala His Tyr Ala Cys Thr Lys
1 5 10

<210> 10
<211> 12
<212> PRT
<213> Artificial Sequence

<220>
<223> hairpin mimetic derived from the general formula Cys-Z-Cys
wherein the alpha amino group of the first amino acid is
acetylated and wherein Z consists of 10 amino acids.

<220>
<221> DISULFID
<222> (1)..(12)

<220>
<221> MOD_RES
<222> (1)..(1)
<223> ACETYLATION

<400> 10

Cys Thr Lys Trp Phe Ser Asn His Tyr Gln Thr Cys
1 5 10

<210> 11
<211> 16
<212> PRT
<213> Artificial Sequence

<220>
<223> Hairpin mimetic derived from the general formula R1-Cys-Z-Cys-R2
wherein the alpha amino group of the first amino acid is
acetylated, wherein Z consists of 10 amino acids, and wherein
both R1 and R2 consist of 2 amino acids.

<220>
<221> MOD_RES
<222> (1)..(1)
<223> ACETYLATION

<220>
<221> DISULFID
<222> (3)..(14)

<400> 11

Glu Thr Cys Thr Lys Trp Phe Ser Asn His Tyr Gln Thr Cys Thr Lys
1 5 10 15

<210> 12
<211> 12
<212> PRT
<213> Artificial Sequence

<220>
<223> Hairpin mimetic derived from the general formula Cys-Z-Cys
wherein the alpha amino group of the first amino acid is
acetylated and wherein Z consists of 10 amino acids.

<220>
<221> DISULFID
<222> (1)..(12)

<220>
<221> MOD_RES
<222> (1)..(1)
<223> ACETYLATION

<400> 12

Cys Thr Lys Trp Phe Leu Ala His Tyr Ala Thr Cys
1 5 10

<210> 13
<211> 16
<212> PRT
<213> Artificial Sequence

<220>
<223> Hairpin mimetic derived from the general formula R1-Cys-Z-Cys-R2
wherein the alpha amino group of the first amino acid is
acetylated, wherein Z consists of 10 amino acids, and wherein
both R1 and R2 consist of 2 amino acids.

<220>
<221> MOD_RES
<222> (1)..(1)
<223> ACETYLATION

<220>
<221> DISULFID
<222> (3)..(14)

<400> 13

Leu Glu Cys Thr Lys Trp Phe Leu Ala His Tyr Ala Thr Cys Lys Val
1 5 10 15

<210> 14
<211> 16
<212> PRT
<213> Artificial Sequence

<220>
<223> Hairpin mimetic derived from the general formula R1-Cys-Z-Cys-R2
wherein the alpha amino group of the first amino acid is
acetylated, wherein Z consists of 10 amino acids, and wherein
both R1 and R2 consist of 2 amino acids.

<220>
<221> MOD_RES
<222> (1)..(1)
<223> ACETYLATION

<220>
<221> DISULFID
<222> (3)..(14)

<400> 14

Asn Gly Cys Thr Lys Trp Phe Leu Ala His Tyr Ala Thr Cys Lys Val
1 5 10 15

<210> 15
<211> 16
<212> PRT
<213> Artificial Sequence

<220>
<223> Hairpin mimetic derived from the general formula R1-Cys-Z-Cys-R2
wherein the alpha amino group of the first amino acid is
acetylated, wherein Z consists of 10 amino acids, and wherein
both R1 and R2 consist of 2 amino acids.

<220>
<221> MOD_RES
<222> (1)..(1)
<223> ACETYLATION

<220>
<221> DISULFID

<222> (3)..(14)

<400> 15

Gly Gly Cys Thr Lys Trp Phe Leu Ala His Tyr Ala Thr Cys Gly Gly
1 5 10 15

<210> 16

<211> 16

<212> PRT

<213> Artificial Sequence

<220>

<223> Hairpin mimetic derived from the general formula R1-Cys-Z-Cys-R2
wherein the alpha amino group of the first amino acid is
acetylated, wherein Z consists of 10 amino acids, and wherein
both R1 and R2 consist of 2 amino acids.

<220>

<221> MOD_RES

<222> (1)..(1)

<223> ACETYLATION

<220>

<221> DISULFID

<222> (3)..(14)

<400> 16

Glu Thr Cys Thr Lys Trp Phe Leu Ala His Tyr Ala Thr Cys Thr Lys
1 5 10 15

<210> 17

<211> 18

<212> PRT

<213> Artificial Sequence

<220>

<223> Hairpin mimetic derived from the general formula R1-Cys-Z-Cys-R2
wherein the alpha amino group of the first amino acid is
acetylated, wherein Z consists of 10 amino acids, and wherein
both R1 and R2 consist of 3 amino acids.

<220>

<221> MOD_RES

<222> (1)..(1)

<223> ACETYLATION

<220>

<221> DISULFID

<222> (4)..(15)

<400> 17

Glu Leu Lys Cys Thr Lys Trp Phe Ser Asn His Tyr Gln Thr Cys Glu
1 5 10 15

Val Lys

<210> 18
<211> 18
<212> PRT
<213> Artificial Sequence

<220>
<223> Hairpin mimetic derived from the general formula R1-Cys-Z-Cys-R2
wherein the alpha amino group of the first amino acid is
acetylated, wherein Z consists of 10 amino acids, and wherein
both R1 and R2 consist of 3 amino acids.

<220>
<221> MOD_RES
<222> (1)..(1)
<223> ACETYLATION

<220>
<221> DISULFID
<222> (4)..(15)

<400> 18

Lys Val Gly Cys Thr Lys Trp Phe Leu Ala His Tyr Ala Thr Cys Gly
1 5 10 15

Leu Glu

<210> 19
<211> 18
<212> PRT
<213> Artificial Sequence

<220>
<223> Hairpin mimetic derived from the general formula R1-Cys-Z-Cys-R2
wherein the alpha amino group of the first amino acid is
acetylated, wherein Z consists of 10 amino acids, and wherein
both R1 and R2 consist of 3 amino acids.

<220>
<221> MOD_RES
<222> (1)..(1)
<223> ACETYLATION

<220>
<221> DISULFID
<222> (4)..(15)

<400> 19

Gly Gly Gly Cys Thr Lys Trp Phe Leu Ala His Tyr Ala Thr Cys Gly
1 5 10 15

Gly Gly

<210> 20
<211> 14
<212> PRT
<213> Artificial Sequence

<220>
<223> Hairpin mimetic derived from the general formula Cys-Z-Cys
wherein the alpha amino group of the first amino acid is
acetylated and wherein Z consists of 12 amino acids.

<220>
<221> DISULFID
<222> (1)..(14)

<220>
<221> MOD_RES
<222> (1)..(1)
<223> ACETYLATION

<400> 20

Cys Gly Thr Lys Trp Phe Ser Asn His Tyr Gln Thr Gly Cys
1 5 10

<210> 21
<211> 18
<212> PRT
<213> Artificial Sequence

<220>
<223> Hairpin mimetic derived from the general formula R1-Cys-Z-Cys-R2
wherein the alpha amino group of the first amino acid is
acetylated, wherein Z consists of 12 amino acids, and wherein
both R1 and R2 consist of 2 amino acids.

<220>
<221> MOD_RES
<222> (1)..(1)
<223> ACETYLATION

<220>
<221> DISULFID
<222> (3)..(16)

<400> 21

Glu Thr Cys Gly Thr Lys Trp Phe Ser Asn His Tyr Gln Thr Gly Cys
1 5 10 15

Thr Lys

<210> 22
<211> 8
<212> PRT
<213> Artificial Sequence

<220>
<223> Core peptide sequence Z taken from the CDR L3 loop of an antibody
described in Jiang, L. et al., Chimia, 2000, 54, 558-563.

<400> 22

Leu Trp Tyr Ser Asn His Trp Val
1 5

<210> 23
<211> 8
<212> PRT
<213> Artificial Sequence

<220>
<223> Modified core peptide sequence Z derived from core peptide
sequence with the SEQ ID NO:22 containing a stabilizing beta-turn
and a beta-sheet sequence according to Chou, P. Y., Fasman, G.
D., J. Mol. Biol., 1977, 115, 135-175.

<400> 23

Lys Trp Phe Ser Asn His Tyr Gln
1 5

<210> 24
<211> 6
<212> PRT
<213> Artificial Sequence

<220>
<223> Core peptide sequence Z constructed from peptide with the SEQ ID
NO:25.

<400> 24

Phe Leu Ala His Tyr Ala

1 5

<210> 25

<211> 10

<212> PRT

<213> Artificial Sequence

<220>

<223> Oligopeptide which does not contain a dedicated stabilizing
beta-turn sequence or a beta-sheet sequence according to Chou, P.
Y., Fasman, G. D., J. Mol. Biol, 1977, 115, 135-175.

<400> 25

Leu Trp Tyr Ser Asn His Trp Val Lys Trp

1 5 10

<210> 26

<211> 39

<212> DNA

<213> Artificial Sequence

<220>

<223> Oligonucleotide No. 1 used to construct insert DNA coding for
template fixed hairpin mimetic of SEQ ID NO:10 and used to
construct insert DNA coding for randomized library template fixed
beta-hairpin mimetics having sequences according to SEQ ID NO:42.

<400> 26

catgcgggg tactttcta ttctcactct gaaacctgc 39

<210> 27

<211> 84

<212> DNA

<213> Artificial Sequence

<220>

<223> Oligonucleotide No. 2 used to construct insert DNA coding for
template fixed hairpin mimetic of SEQ ID NO:10.

<400> 27

catgttcgg ccgagccacc accttggtg caggtctgat aatggttgct gaaccatttg 60

gtgcagggtt cagagtgaga atag

84

<210> 28

<211> 30

<212> DNA

<213> Artificial Sequence

<220>

<223> DNA sequence coding for the peptide shown in SEQ ID NO:8.

<400> 28
tgcaaatgg tccctggcgca ttatgcgtgc 30

<210> 29
<211> 42
<212> DNA
<213> Artificial Sequence

<220>
<223> DNA sequence coding for the peptide shown in SEQ ID NO:9.

<400> 29
gaaacctgca aatggttcct ggccgcattat gcgtgcacca aa 42

<210> 30
<211> 36
<212> DNA
<213> Artificial Sequence

<220>
<223> DNA sequence coding for the peptide shown in SEQ ID NO:10.

<400> 30
tgcaccaat ggttcagcaa ccattatcag acctgc 36

<210> 31
<211> 48
<212> DNA
<213> Artificial Sequence

<220>
<223> DNA sequence coding for the peptide shown in SEQ ID NO:11.

<400> 31
gaaacctgca ccaaatgg t cagcaaccat tatcagacct gcaccaaa 48

<210> 32
<211> 36
<212> DNA
<213> Artificial Sequence

<220>
<223> DNA sequence coding for the peptide shown in SEQ ID NO:12.

<400> 32
tgcaccaat ggttcctggc gcattatgcg acctgc 36

<210> 33
<211> 48
<212> DNA
<213> Artificial Sequence

<220>
<223> DNA sequence coding for the peptide shown in SEQ ID NO:13.

<400> 33
cttggatgca ccaaattgggtt cctggcgcat tatgcgaccc gcaaaagtt 48

<210> 34
<211> 48
<212> DNA
<213> Artificial Sequence

<220>
<223> DNA sequence coding for the peptide shown in SEQ ID NO:14.

<400> 34
aacgggttgca ccaaattgggtt cctggcgcat tatgcgaccc gcaaaagtt 48

<210> 35
<211> 48
<212> DNA
<213> Artificial Sequence

<220>
<223> DNA sequence coding for the peptide shown in SEQ ID NO:15.

<400> 35
gggtgggttgca ccaaattgggtt cctggcgcat tatgcgaccc ggggggggt 48

<210> 36
<211> 48
<212> DNA
<213> Artificial Sequence

<220>
<223> DNA sequence coding for the peptide shown in SEQ ID NO:16.

<400> 36
gaaacacctgca ccaaattgggtt cctggcgcat tatgcgaccc gcacccaaa 48

<210> 37
<211> 54
<212> DNA
<213> Artificial Sequence

<220>
<223> DNA sequence coding for the peptide shown in SEQ ID NO:17.

<400> 37
gaactgaaat gcacccaaatg gttcagcaac cattatcaga cctgcgaagt taaa 54

<210> 38

<211> 54
<212> DNA
<213> Artificial Sequence

<220>

<223> DNA sequence coding for the peptide shown in SEQ ID NO:18.

<400> 38

aaagttggtt gcaccaaatg gttcctggcg cattatgcga cctgcggctt ggaa 54

<210> 39
<211> 54
<212> DNA
<213> Artificial Sequence

<220>

<223> DNA sequence coding for the peptide shown in SEQ ID NO:19.

<400> 39

ggtgtggctt gcaccaaatg gttcctggcg cattatgcga cctgcggcg gggt 54

<210> 40
<211> 42
<212> DNA
<213> Artificial Sequence

<220>

<223> DNA sequence coding for the peptide shown in SEQ ID NO:20.

<400> 40

tgcgttacca aatggttcag caaccattat cagaccggtt gc 42

<210> 41
<211> 54
<212> DNA
<213> Artificial Sequence

<220>

<223> DNA sequence coding for the peptide shown in SEQ ID NO:21.

<400> 41

gaaacctgcg gtaccaaatg gttcagcaac cattatcaga ccggttgcac caaa 54

<210> 42
<211> 48
<212> DNA
<213> Artificial Sequence

<220>

<223> DNA sequence of randomized template fixed beta-hairpin mimetic Phage library.

```

<220>
<221> misc_feature
<222> (10)..(11)
<223> n is a, c, g, or t

<220>
<221> misc_feature
<222> (13)..(14)
<223> n is a, c, g, or t

<220>
<221> misc_feature
<222> (16)..(17)
<223> n is a, c, g, or t

<220>
<221> misc_feature
<222> (28)..(29)
<223> n is a, c, g, or t

<220>
<221> misc_feature
<222> (31)..(32)
<223> n is a, c, g, or t

<220>
<221> misc_feature
<222> (34)..(35)
<223> n is a, c, g, or t

<220>
<221> misc_feature
<222> (37)..(38)
<223> n is a, c, g, or t

<400> 42
gaaacctgcn nknnknnkcg tggtgacnnk nnknnknnkt gcaccaaa

```

48

```

<210> 43
<211> 16
<212> PRT
<213> Artificial Sequence

<220>
<223> Translated protein sequence of a randomized template fixed
      beta-hairpin mimetic phage library

<220>
<221> DISULFID
<222> (3)..(14)

<220>
<221> MISC_FEATURE
<222> (4)..(6)
<223> Xaa can be any naturally occurring amino acid

```

<220>
<221> MISC_FEATURE
<222> (10)..(13)
<223> Xaa can be any naturally occurring amino acid

<400> 43

Glu Thr Cys Xaa Xaa Xaa Arg Gly Asp Xaa Xaa Xaa Xaa Cys Thr Lys
1 5 10 15

<210> 44
<211> 84
<212> DNA
<213> Artificial Sequence

<220>
<223> Oligonucleotide No. 3 used to construct insert DNA coding for
randomized library template fixed beta-hairpin mimetics having
sequences according to SEQ ID NO:42.

<220>
<221> misc_feature
<222> (34)..(35)
<223> n is a, c, g, or t

<220>
<221> misc_feature
<222> (37)..(38)
<223> n is a, c, g, or t

<220>
<221> misc_feature
<222> (40)..(41)
<223> n is a, c, g, or t

<220>
<221> misc_feature
<222> (43)..(44)
<223> n is a, c, g, or t

<220>
<221> misc_feature
<222> (55)..(56)
<223> n is a, c, g, or t